#### Certificate of Mailing/Transmission (37 C.F.R. § 1.8(a)):

[X] Pursuant to 37 C.F.R. § 1.8, I hereby certify that this paper and all enclosures are being deposited with the United States Postal Service as first class mail on the date indicated below in an envelope addressed to the Assistant Commissioner for Patents, Washington D.C. 20231.

Dated: Mysel 6, 2001

Name of Person Certifying:

## Printed Name: Jocel

#### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant:

Oxana Ibraghimov-Beskrovnaya et al.

Examiner:

Not Yet Assigned

Filing Date:

April 26, 2001

Serial No.:

09/830,506

Group Art Unit: Not Yet Assigned

Title:

COMPOSITIONS AND METHODS FOR TREATING POLYCYSTIC KIDNEY DISEASE

### STATEMENT REGARDING SEQUENCE LISTING UNDER 37 C.F.R.§§ 1.821-1.825

Commissioner for Patents Washington, D.C. 20231

Dear Sir:

The Applicant hereby declares that the content of the computer-readable copy of the Sequence Listing furnished herewith is the same as the written copy of the Sequence Listing.

Respectfully submitted,

Antoinette F. Konski Registration No. 34,202

Mailing Address:

McCutchen, Doyle, Brown & Enersen, LLP

Three Embarcadero Center, Suite 1800

San Francisco, CA 94111 Telephone: (650) 849-4936

Facsimile: (650) 849-4800

· . . .

# PTO/PCT Rec'd 10 AUG 2001

SEQUENCE LISTING

```
<110> Genzyme Corporation
<120> COMPOSITIONS AND METHODS FOR IDENTIFYING
 AND TREATING POLYCYSTIC KIDNEY DISEASE
<130> 126881206140
<140> PCT/US99/
<141> 1999-10-22
<150> 60/105,731
<151> 1998-10-28
<150> 60/105,876
<151> 1998-10-27
<150> 60/141,175
<151> 1999-06-25
<160> 3
<170> FastSEQ for Windows Version 3.0
<210> 1
<211> 14060
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> (135)...(13040)
<221> misc feature
<222> (2621)...(2710)
<223> epitope in the loop region of the polycystin
       transmembrane domain
<221> misc feature
<222> (2734)...(3094)
<223> epitope in the loop region of the polycystin
       transmembrane domain
<221> misc feature
 <222> (3166)...(3300)
 <223> epitope in the loop region of the polycystin
       transmembrane domain
 <221> misc feature
 \langle 222 \rangle (336\overline{4}) \dots (3578)
 <223> epitope in the loop region of the polycystin
       transmembrane domain
 <221> misc feature
 \langle 222 \rangle (362\overline{3}) \dots (3688)
 <223> epitope in the loop region of the polycystin
       transmembrane domain
 <221> misc_feature
 <222> (3710)...(3914)
 <223> epitope in the loop region of the polycystin
        transmembrane domain
 <221> misc_feature
```

PCT/US99/25091

<222> (3931)...(4046) <223> epitope in the loop region of the polycystin transmembrane domain <221> misc feature  $\langle 222 \rangle$   $(216\overline{6}) \dots (2599)$ <223> epitope outside the loop region but within the polycystin transmembrane domain <221> misc\_feature <222> (4097)...(4302) <223> epitope in the loop region of the polycystin transmembrane domain <221> misc feature  $\langle 222 \rangle (414\overline{8}) \dots (4219)$ <223> epitope in the loop region of the polycystin transmembrane domain <221> misc\_feature <222> (4220)...(4302) <223> epitope in the loop region of the polycystin transmembrane domain <221> misc feature <222> (27)...(360) <223> epitope in the loop region of the polycystin transmembrane domain <221> misc\_feature <222> (843)...(1200) <223> Ig-like domain of polycystin <221> misc feature  $\langle 222 \rangle$   $(120\overline{5}) \dots (1625)$ <223> Ig-like domain of polycystin <221> misc\_feature <222> (1626)...(2136) <223> Ig-like domain of polycystin <221> misc feature  $\langle 222 \rangle$   $(216\overline{6}) \dots (2599)$ <223> isolated polypeptide <400> 1 60 gctcagcage aggtcgcggc cgcagcccca tccagcccgc gcccgccatg ccgtccgcgg 120 geoecgectg agetgeggte teegegegeg ggegggeetg gggaeggegg ggeeatgege 170 gegetgeect aacg atg eeg eee gee geg eee gee etg geg etg gee Met Pro Pro Ala Ala Pro Ala Arg Leu Ala Leu Ala 218 Leu Gly Leu Gly Leu Trp Leu Gly Ala Leu Ala Gly Gly Pro Gly Arg 266 gge tge ggg eec tge gag eec eec tge etc tge gge eea geg eec gge Gly Cys Gly Pro Cys Glu Pro Pro Cys Leu Cys Gly Pro Ala Pro Gly 314 ged ged tgd ege gtd aad tgd teg ggd egg ggg etg egg acg etd ggt Ala Ala Cys Arg Val Asn Cys Ser Gly Arg Gly Leu Arg Thr Leu Gly 50 362 ccc gcg ctg cgc atc ccc gcg gac gcc aca gcg cta gac gtc tcc cac Pro Ala Leu Arg Ile Pro Ala Asp Ala Thr Ala Leu Asp Val Ser His

								3									
				65					70					75		-	-
aac Asn	ctg Leu	ctc Leu	cgg Arg 80	gcg Ala	ctg Leu	gac Asp	gtt Val	ggg Gly 85	ctc Leu	ctg Leu	gcg Ala	aac Asn	ctc Leu 90	tcg Ser	gcg Ala	4.3	10
ctg Leu	gca Ala	gag Glu 95	ctg Leu	gat Asp	ata Ile	agc Ser	aac Asn 100	aac Asn	aag Lys	att Ile	tct Ser	acg Thr 105	tta Leu	gaa Glu	gaa Glu	4 5	58
gga Gly	ata Ile 110	ttt Phe	gct Ala	aat Asn	tta Leu	ttt Phe 115	aat Asn	tta Leu	agt Ser	gaa Glu	ata Ile 120	aac Asn	ctg Leu	agt Ser	GJ À àdà	50	06
aac Asn 125	ccg Pro	ttt Phe	gag Glu	tgt Cys	gac Asp 130	tgt Cys	ggc Gly	ctg Leu	gcg Ala	tgg Trp 135	ctg Leu	ccg Pro	cga Arg	tgg Trp	gcg Ala 140	5	54
gag Glu	gag Glu	cag Gln	cag Gln	gtg Val 145	cgg Arg	gtg Val	gtg Val	cag Gln	ccc Pro 150	gag Glu	gca Ala	gcc Ala	acg Thr	tgt Cys 155	gct Ala	6	02
Gly	cct Pro	ggc Gly	tcc Ser 160	ctg Leu	gct Ala	ggc Gly	cag Gln	cct Pro 165	ctg Leu	ctt Leu	ggc Gly	atc Ile	ccc Pro 170	ttg Leu	ctg Leu	6	50
gac Asp	agt Ser	ggc Gly 175	tgt Cys	ggt Gly	gag Glu	gag Glu	tat Tyr 180	gtc Val	gcc Ala	tgc Cys	ctc Leu	cct Pro 185	gac Asp	aac Asn	agc Ser	6	598
tca Ser	ggc Gly 190	acc Thr	gtg Val	gca Ala	gca Ala	gtg Val 195	tcc Ser	ttt Phe	tca Ser	gct Ala	gcc Ala 200	cac His	gaa Glu	ggc Gly	ctg Leu	7	46
ctt Leu 205	cag Gln	cca Pro	gag Glu	gcc Ala	tgc Cys 210	agc Ser	gcc Ala	ttc Phe	tgc Cys	ttc Phe 215	Ser	acc Thr	ggc Gly	cag Gln	ggc Gly 220	7	794
ctc Leu	gca Ala	gcc Ala	ctc Leu	tcg Ser 225	gag Glu	cag Gln	ggc Gly	tgg Trp	tgc Cys 230	Leu	tgt Cys	ggg Gly	gcg Ala	gcc Ala 235	Gin		842
ccc Pro	tcc Ser	agt Ser	gcc Ala 240	Ser	ttt Phe	gcc Ala	tgc Cys	ctg Leu 245	Ser	cto Leu	tgo Cys	tcc Ser	ggc Gly 250	Pro	ccg Pro	\$	890
cca Pro	cct Pro	cct Pro 255	Ala	ccc Pro	acc Thr	tgt Cys	agg Arg 260	, Gly	Pro	aco Thi	c cto Lev	cto Lev 265	ı Glm	cac His	gtc Val		938
ttc Phe	cct Pro 270	Ala	tcc Ser	cca Pro	ggg	gcc Ala 275	Thr	ctg Leu	gto Val	g ggg	g cco y Pro 280	o His	gga Gly	cct Pro	ctg Leu		986
gcc Ala 285	Ser	ggc Gly	caç Glr	cta Leu	gca Ala 290	Alā	tto Phe	c cac e His	ato Ile	c gc e Ala 29	a Al	c ccc a Pro	g cto Lev	c cct i Pro	gtc Val 300	1	034
act Thr	gcc Ala	aca Thi	cgc Arg	tgg Trp 305	Asp	tto Phe	gga Gly	a gad y Asp	gg Gl; 31	y Se	c gc	c gad a Gl	g gto u Val	g gat l As <sub>l</sub> 31:	t gcc p Ala 5	1	.082
gct Ala	ggç Gl	g ccq y Pro	g gct o Ala 320	a Ala	tco a Ser	g cat	cgo s Āro	g Ty:	c Va	g ct l Le	g cc u Pr	t gg o Gl	g cgo y Aro 33	д Ту	t cac r His	1	130

								4								
gtg Val	acg Thr	gcc Ala 335	gtg Val	ctg Leu	gcc Ala	Leu (	999 Gly 340	gcc Ala	ggc Gly	tca Ser	gcc Ala	ctg Leu 345	ctg Leu	GJ À ààà	aca Thr	1178
gac Asp	gtg Val 350	cag Gln	gtg Val	gaa Glu	Ala	gca Ala 355	cct Pro	gcc Ala	gcc Ala	ctg Leu	gag Glu 360	ctc Leu	gtg Val	tgc Cys	ccg Pro	1226
tcc Ser 365	tcg Ser	gtg Val	cag Gln	agt Ser	gac Asp 370	gag Glu	agc Ser	ctt Leu	gac Asp	ctc Leu 375	agc Ser	atc Ile	cag Gln	aac Asn	cgc Arg 380	1274
ggt Gly	ggt Gly	tca Ser	ggc Gly	ctg Leu 385	gag Glu	gcc Ala	gcc Ala	tac Tyr	agc Ser 390	atc Ile	gtg Val	gcc Ala	ctg Leu	ggc Gly 395	gag Glu	1322
gag Glu	ccg Pro	gcc Ala	cga Arg 400	gcg Ala	gtg Val	cac His	ccg Pro	ctc Leu 405	tgc Cys	ccc Pro	tcg Ser	gac Asp	acg Thr 410	gag Glu	atc Ile	1370
ttc Phe	cct Pro	ggc Gly 415	aac Asn	ggg Gly	cac His	tgc Cys	tac Tyr 420	cgc Arg	ctg Leu	gtg Val	gtg Val	gag Glu 425	aag Lys	gcg Ala	gcc Ala	1418
tgg Trp	ctg Leu 430	cag Gln	gcg Ala	cag Gln	gag Glu	cag Gln 435	tgt Cys	cag Gln	gcc Ala	tgg Trp	gcc Ala 440	Gly	gcc Ala	gcc Ala	ctg Leu	1466
gca Ala 445	atg Met	gtg Val	gac Asp	agt Ser	ccc Pro 450	gcc Ala	gtg Val	cag Gln	cgc Arg	ttc Phe 455	ctg Leu	gtc Val	tcc Ser	cgg Arg	gtc Val 460	1514
acc Thr	agg Arg	agc Ser	cta Leu	gac Asp 465	gtg Val	tgg Trp	atc Ile	ggc Gly	ttc Phe 470	tcg Ser	act Thr	gtg Val	cag Gln	ggg Gly 475	Val	1562
gag Glu	gtg Val	ggc Gly	cca Pro 480	gcg Ala	ccg Pro	cag Gln	ggc Gly	gag Glu 485	gcc Ala	ttc Phe	ago Ser	ctg Leu	gag Glu 490	Ser	tgc Cys	1610
cag Gln	aac Asn	tgg Trp 495	Leu	ccc Pro	Gly ggg	gag Glu	cca Pro 500	His	cca Pro	gcc Ala	aca Thr	gcc Ala 505	GIU	cac His	tgc Cys	1658
gtc Val	cgg Arg 510	Let	ggg Gly	ccc Pro	acc Thr	ggg Gly 515	tgg Trp	tgt Cys	aac Asn	acc Thr	gad Asp 520	) Let	tgc Cys	tca Sei	a gcg Ala	1706
ccg Pro 525	His	ago Ser	tac Tyr	gtc Val	tgc Cys 530	Glu	ctg Leu	cag Gln	cco Pro	gga Gly 535	/ Gl	c cca y Pro	a gto Val	g caq L Gli	g gat n Asp 540	1754
gcc Ala	gaç Glu	g aad 1 Asr	c ctc n Leu	cto Leu 545	ı Val	gga Gly	gcc	g ccc Pro	agt Ser 550	Gly	g gad y Asj	c cto p Le	g caq ı Glı	g gga n Gl; 55	a ccc y Pro 5	1802
cto Lei	g acq ı Thi	g cct	t cto 560	ı Ala	cag Gln	rcag Gln	gad Asp	ggc Gly 569	/ Le	tca 1 Sei	a gc r Al	c cc a Pr	g cae 5 Hi:	S GI	g ccc u Pro	1850
gto Vai	g gad l Gli	g gte u Va 57	l Met	g gta Val	a tto L Phe	ccg Pro	999 6 Gly 580	y Lei	g cg	cto g Le	g ag u Se	c cg r Ar 58	g GI	a gc u Al	c ttc a Phe	1898
ct. Le	c ac	c ac r Th	g gco r Ala	c gaa a Glu	a ttt u Phe	ggç Gl	ace Th	c cad	g ga n Gl	g ct u Le	c cg u Ar	g cg g Ar	g cc g Pr	c gc o Al	c cag a Gln	1946

								5								
	590					595					600					-
ctg Leu 605	cgg Arg	ctg Leu	cag Gln	gtg Val	tac Tyr 610	cgg Arg	ctc Leu	ctc Leu	agc Ser	aca Thr 615	gca Ala	ggg Gly	acc Thr	ccg Pro	gag Glu 620	1994
												agg Arg				2042
gcc Ala	ccc Pro	gcg Ala	tgc Cys 640	atg Met	cca Pro	G] À ddd	gga Gly	cgc Arg 645	tgg Trp	tgc Cys	cct Pro	gga Gly	gcc Ala 650	aac Asn	atc Ile	2090
tgc Cys	ttg Leu	ccg Pro 655	ctg Leu	gac Asp	gcc Ala	tct Ser	tgc Cys 660	cac His	ccc Pro	cag Gln	gcc Ala	tgc Cys 665	gcc Ala	aat Asn	ggc Gly	2138
tgc Cys	acg Thr 670	tca Ser	ggg Gly	cca Pro	ggg Gly	cta Leu 675	ccc Pro	ggg Gly	gcc Ala	ccc Pro	tat Tyr 680	gcg Ala	cta Leu	tgg Trp	aga Arg	2186
gag Glu 685	ttc Phe	ctc Leu	ttc Phe	tcc Ser	gtt Val 690	gcc Ala	gcg Ala	ggg Gly	ccc Pro	ccc Pro 695	gcg Ala	cag Gln	tac Tyr	tcg Ser	gtc Val 700	2234
acc Thr	ctc Leu	cac His	ggc Gly	cag Gln 705	gat Asp	gtc Val	ctc Leu	atg Met	ctc Leu 710	cct Pro	ggt Gly	gac Asp	ctc Leu	gtt Val 715	ggc Gly	2282
ttg Leu	cag Gln	cac His	gac Asp 720	gct Ala	ggc Gly	cct Pro	ggc Gly	gcc Ala 725	ctc Leu	ctg Leu	cac His	tgc Cys	tcg Ser 730	ccg Pro	gct Ala	2330
ccc Pro	ggc Gly	cac His 735	cct Pro	ggt Gly	ccc Pro	cag Gln	gcc Ala 740	ccg Pro	tac Tyr	ctc Leu	tcc Ser	gcc Ala 745	aac Asn	gcc Ala	tcg Ser	2378
tca Ser	tgg Trp 750	ctg Leu	ccc Pro	cac His	ttg Leu	cca Pro 755	gcc Ala	cag Gln	ctg Leu	gag Glu	ggc Gly 760	act Thr	tgg Trp	gcc Ala	tgc Cys	2426
cct Pro 765	gcc Ala	tgt Cys	gcc Ala	ctg Leu	cgg Arg 770	ctg Leu	ctt Leu	gca Ala	gcc Ala	acg Thr 775	Glu	cag Gln	ctc Leu	acc Thr	gtg Val 780	2474
ctg Leu	ctg Leu	ggc Gly	ttg Leu	agg Arg 785	ccc Pro	aac Asn	cct Pro	gga Gly	ctg Leu 790	Arg	atg Met	cct Pro	ggg Gly	cgc Arg 795	Tyr	2522
gag Glu	gtc Val	cgg Arg	gca Ala 800	gag Glu	gtg Val	ggc Gly	aat Asn	ggc Gly 805	Val	tcc Ser	agg Arg	cac His	aac Asn 810	Leu	tcc Ser	2570
tgc Cys	agc Ser	ttt Phe 815	Asp	gtg Val	gtc Val	tcc Ser	cca Pro 820	Val	gct Ala	ggg	cto Leu	cgg Arg 825	Val	ato Ile	tac Tyr	2618
cct Pro	gcc Ala 830	Pro	cgc Arg	gac Asp	ggc Gly	cgc Arg 835	Leu	tac Tyr	gtç Val	Pro	acc Thr 840	Asn	ggc Gly	tca Ser	gcc Ala	2666
ttg Leu 845	ı Val	cto Lev	cag Gln	gtg Val	gac Asp 850	Ser	ggt	gcc Ala	aac Asr	gco Ala 855	a Thi	g gcc Ala	aco Thr	g gct Ala	cgc Arg 860	2714

WO 00/24780	6	PCT/US99/25091
	<b>41</b>	

•	<b>VO</b> 0	0/247	80						_						PC	CT/US99/	25091
									6								_
t T	gg Trp	cct Pro	Gly Ggg	ggc Gly	agt Ser 865	gtc Val	agc Ser	gcc Ala	cgc Arg	ttt Phe 870	gag Glu	aat Asn	gtc Val	tgc Cys	cct Pro 875	gcc Ala	2762
															gat Asp		2810
															cac His		2858
7	Val	gac Asp 910	gtg Val	gtg Val	gtg Val	gaa Glu	aac Asn 915	agc Ser	gcc Ala	agc Ser	cgg Arg	gcc Ala 920	aac Asn	ctc Leu	agc Ser	ctg Leu	2906
i	egg Arg 925	Val	acg Thr	gcg Ala	gag Glu	gag Glu 930	ccc Pro	atc Ile	tgt Cys	ggc Gly	ctc Leu 935	cgc Arg	gcc Ala	acg Thr	ccc Pro	agc Ser 940	2954
•	ccc Pro	gag Glu	gcc Ala	cgt Arg	gta Val 945	ctg Leu	cag Gln	gga Gly	gtc Val	cta Leu 950	gtg Val	agg Arg	tac Tyr	agc Ser	ccc Pro 955	gtg Val	3002
	gtg Val	gag Glu	gcc Ala	ggc Gly 960	tcg Ser	gac Asp	atg Met	gtc Val	ttc Phe 965	cgg Arg	tgg Trp	acc Thr	atc Ile	aac Asn 970	gac Asp	aag Lys	3050
	cag Gln	tcc Ser	ctg Leu 975	acc Thr	ttc Phe	cag Gln	aac Asn	gtg Val 980	gtc Val	ttc Phe	aat Asn	gtc Val	att Ile 985	tat Tyr	cag Gln	agc Ser	3098
	gcg Ala	gcg Ala 990	Val	ttc Phe	aag Lys	ctc Leu	tca Ser 99	Leu	acg Thr	gcc Ala	tcc Ser	aac Asn 100	His	gtg Val	agc Ser	aac Asn	3146
	gtc Val 1005	Thr	gtg Val	aac Asn	tac Tyr	aac Asn 101	Val	acc Thr	gtg Val	gag Glu	cgg Arg 101	Met	aac Asn	agg Arg	atg Met	cag Gln 1020	3194
	ggt Gly	ctg Leu	cag Gln	gtc Val	tcc Ser 102	Thr	gtg Val	ccg Pro	gcc Ala	gtg Val 103	Leu	tcc Ser	ccc Pro	aat Asn	gcc Ala 103	Thr	3242
	cta Leu	gca Ala	ctg Leu	acg Thr 104	Ala	ggc Gly	gtg Val	ctg Leu	gtg Val 104	Asp	tcg Ser	gcc Ala	gtg Val	gag Glu 105	gtg Val 0	gcc Ala	3290
	ttc Phe	ctg Leu	tgg Trp 105	Asn	ttt Phe	ggg	gat Asp	ggg Gly 106	Glu	cag Gln	gcc Ala	ctc Leu	cac His	Gln	ttc Phe	cag Gln	3338
	cct Pro	ccg Pro 107	Tyr	aac Asn	gag Glu	tcc	ttc Phe 107	Pro	gtt Val	cca Pro	gac Asp	ccc Pro 108	Ser	g gtg Val	g gcc . Ala	cag Gln	3386
	gtg Val 108	Leu	gtg Val	gag Glu	cac His	aat Asn 109	. Val	ato Met	g cac His	acc Thr	tac Tyr 109	Ala	gco Ala	c cca a Pro	a ggt o Gly	gag Glu 1100	3434
	tac Tyr	ctc Leu	cto Leu	acc Thr	gtg Val	. Let	gca Ala	a tct a Sei	aat Asr	gco Ala 111	a Ph∈	gaç Glu	g aad n Asi	c cto	g acq ı Thr 111	g cag Gln 5	3482
	cag	gto	g cct	gtg Val	ago	gtç	g cgc	geo a Ala	t tco	c cto	g ccc	tco Sei	gto Vai	g gct l Ala	t gto a Val	g ggt L Glv	3530

	7		1 0 17 0 0 7 7 7 2 .	
1120	11	125	1130	-
gtg agt gac ggc g Val Ser Asp Gly V 1135	tc ctg gtg gcc gg al Leu Val Ala Gl 1140	ge egg eee gte ace ly Arg Pro Val Thr 1145	Phe Tyr Pro	3578
cac ccg ctg ccc t His Pro Leu Pro S 1150	cg cct ggg ggt gt er Pro Gly Gly Va 1155	tt ctt tac acg tgg al Leu Tyr Thr Trp 1160	gac ttc ggg Asp Phe Gly	3626
gac ggc tcc cct g Asp Gly Ser Pro V 1165	tc ctg acc cag ac al Leu Thr Gln Se 1170	gc cag ccg gct gcc er Gln Pro Ala Ala 1175	aac cac acc Asn His Thr 1180	3674
Tyr Ala Ser Arg G	gc acc tac cac gt ly Thr Tyr His Va 185	tg cgc ctg gag gtc al Arg Leu Glu Val 1190	aac aac acg Asn Asn Thr 1195	3722
gtg agc ggt gcg g Val Ser Gly Ala A 1200	lla Ala Gln Ala A	at gtg cgc gtc ttt sp Val Arg Val Phe 205	gag gag ctc Glu Glu Leu 1210	3770
cgc gga ctc agc c Arg Gly Leu Ser V 1215	rtg gac atg agc c Val Asp Met Ser L 1220	tg gcc gtg gag cag eu Ala Val Glu Gln 122	Gly Ala Pro	3818
gtg gtg gtc agc g Val Val Val Ser A 1230	gcc gcg gtg cag a Ala Ala Val Gln T 1235	cg ggc gac aac atc Thr Gly Asp Asn Ile 1240	acg tgg acc Thr Trp Thr	3866
ttc gac atg ggg o Phe Asp Met Gly <i>I</i> 1245	gac ggc acc gtg c Asp Gly Thr Val L 1250	tg tcg ggc ccg gag Leu Ser Gly Pro Glu 1255	gca aca gtg Ala Thr Val 1260	391 ·
Glu His Val Tyr	ctg cgg gca cag a Leu Arg Ala Gln A 1265	aac tgc aca gtg acc Asn Cys Thr Val Thr 1270	gtg ggt gcg Val Gly Ala 1275	3962
gcc agc ccc gcc Ala Ser Pro Ala 1280	Gly His Leu Ala A	cgg agc ctg cac gtg Arg Ser Leu His Val 1285	ctg gtc ttc Leu Val Phe 1290	4010
gtc ctg gag gtg Val Leu Glu Val 1295	ctg cgc gtt gaa c Leu Arg Val Glu F 1300	ccc gcc gcc tgc ato Pro Ala Ala Cys Ile 130	Pro Thr Gin	4058
cct gac gcg cgg Pro Asp Ala Arg 1310	ctc acg gcc tac o Leu Thr Ala Tyr v 1315	gtc acc ggg aac ccc Val Thr Gly Asn Pro 1320	g gcc cac tac o Ala His Tyr	4106
ctc ttc gac tgg Leu Phe Asp Trp 1325	acc ttc ggg gat of Thr Phe Gly Asp of 1330	ggc tcc tcc aac acc Gly Ser Ser Asn Th: 1335	g acc gtg cgg r Thr Val Arg 1340	4154
ggg tgc ccg acg Gly Cys Pro Thr	gtg aca cac aac Val Thr His Asn 1345	ttc acg cgg agc gg Phe Thr Arg Ser Gl 1350	c acg ttc ccc y Thr Phe Pro 1355	4202
ctg gcg ctg gtg Leu Ala Leu Val 1360	Leu Ser Ser Arg	gtg aac agg gcg ca Val Asn Arg Ala Hi 1365	t tac ttc acc s Tyr Phe Thr 1370	4250
agc atc tgc gtg Ser Ile Cys Val 1375	gag cca gag gtg Glu Pro Glu Val 1380	ggc aac gtc acc ct Gly Asn Val Thr Le ) 13	g cag cca gag u Gln Pro Glu 85	4298

								8						-		
agg Arg	cag Gln 1390	Phe	gtg Val	cag Gln	ctc Leu	ggg Gly 1395	Asp	gag Glu	gcc Ala	tgg Trp	ctg Leu 1400	Val	gca Ala	tgt Cys	gcc Ala	4346
tgg Trp 1405	Pro	ccg Pro	ttc Phe	ccc Pro	tac Tyr 1410	Arg	tac Tyr	acc Thr	tgg Trp	gac Asp 1415	ttt Phe	ggc Gly	acc Thr	gag Glu	gaa Glu 1420	4394
					Ala					Val	acg Thr				Arg	4442
				Tyr					Thr		tcc Ser			Ile		4490
			Asp					Glu			gag Glu		Val			4538
		Ile					Ser				gag Glu 1480	Leu				4586
tac Tyr 1485	Leu	ttc Phe	tct Ser	gct Ala	gtg Val 1490	Gly	cgt Arg	ggg Gly	cgc Arg	ccc Pro 1495	gcc Ala	agc Ser	tac Tyr	ctg Leu	tgg Trp 1500	4634
					Gly					Pro	gag Glu				Āla	4682
				Gly					Arg		gcc Ala			Asn		4730
			Ser					Asn			gtg Val		Arg			4778
		Leu					Ser				gtg Val 1560	Pro				4826
	Val					Ser					agt Ser 5					4874
					Asp					Ile	cct Pro				Thr	4922
				Phe					Thr		aat Asn			Val		4970
			Glu					Gln			atc Ile		Val			5018
		Leu					Gln				ggt Gly 164	Gly			ttc Phe	5066
															acc	5114

	9		
1645	1650	1655	1660
aac gtc tcc tac agc	Trp Thr Ala Trp	agg gac agg ggc ccg	gcc ctg 5162
Asn Val Ser Tyr Ser		Arg Asp Arg Gly Pro	Ala Leu
166		1670	1675
gcc ggc agc ggc aaa	ggc ttc tcg ctc	acc gtg ctc gag gcc	Gly Thr
Ala Gly Ser Gly Lys	Gly Phe Ser Leu	Thr Val Leu Glu Ala	
1680	168	5 1690	
tac cat gtg cag ctg Tyr His Val Gln Leu 1695	cgg gcc acc aac Arg Ala Thr Asn 1700	atg ctg ggc agc gcc Met Leu Gly Ser Ala 1705	tgg gcc 5258 Trp Ala
gac tgc acc atg gac Asp Cys Thr Met Asp 1710	ttc gtg gag cct Phe Val Glu Pro 1715	gtg ggg tgg ctg atg Val Gly Trp Leu Met 1720	gtg acc 5306 Val Thr
gcc tcc ccg aac cca	gct gcc gtc aac	aca agc gtc acc ctc	agt gcc 5354
Ala Ser Pro Asn Pro	Ala Ala Val Asn	Thr Ser Val Thr Leu	Ser Ala
1725	1730	1735	1740
gag ctg gct ggt ggc	Ser Gly Val Val	tac act tgg tcc ttg	gag gag 5402
Glu Leu Ala Gly Gly		Tyr Thr Trp Ser Leu	Glu Glu
174		1750	1755
ggg ctg agc tgg gag	acc tcc gag cca	ttt acc acc cat agc	Phe Pro
Gly Leu Ser Trp Glu	Thr Ser Glu Pro	Phe Thr Thr His Ser	
1760	176	5 1770	
aca ccc ggc ctg cac Thr Pro Gly Leu His 1775	ttg gtc acc atg Leu Val Thr Met 1780	acg gca ggg aac ccg Thr Ala Gly Asn Pro 1785	ctg ggc 5498 Leu Gly
tca gcc aac gcc acc Ser Ala Asn Ala Thr 1790	gtg gaa gtg gat Val Glu Val Asp 1795	gtg cag gtg cct gtg Val Gln Val Pro Val 1800	agt ggc 5546 Ser Gly
ctc agc atc agg gcc	agc gag ccc gga	ggc agc ttc gtg gcg	gcc ggg 5594
Leu Ser Ile Arg Ala	Ser Glu Pro Gly	Gly Ser Phe Val Ala	Ala Gly
1805	1810	1815	1820
tcc tct gtg ccc ttt	Trp Gly Gln Leu	gcc acg ggc acc aat	gtg agc 5642
Ser Ser Val Pro Phe		Ala Thr Gly Thr Asn	Val Ser
182		1830	1835
		agc aag cgt ggc cct Ser Lys Arg Gly Pro 5 1850	His Val
acc atg gtc ttc ccg Thr Met Val Phe Pro 1855	gat gct ggc acc Asp Ala Gly Thr 1860	ttc tcc atc cgg ctc Phe Ser Ile Arg Leu 1865	aat gcc 5738 Asn Ala
tcc aac gca gtc agc Ser Asn Ala Val Ser 1870	tgg gtc tca gcc Trp Val Ser Ala 1875	acg tac aac ctc acg Thr Tyr Asn Leu Thr 1880	gcg gag 5786 Ala Glu
gag ccc atc gtg ggc	ctg gtg ctg tgg	gcc agc agc aag gtg	gtg gcg 5834
Glu Pro Ile Val Gly	Leu Val Leu Trp	Ala Ser Ser Lys Val	Val Ala
.1885	1890	1895	1900
ccc ggg cag ctg gtc	His Phe Gln Ile	ctg ctg gct gcc ggc	tca gct 5882
Pro Gly Gln Leu Val		Leu Leu Ala Ala Gly	Ser Ala
190		1910	1915

110 00/24/80	10	PC1/US99/25091
gtc acc ttc cgc ct	g cag gtc ggc ggg gcc a	aac ccc gag gtg ctc ccc 5930
Val Thr Phe Arg Le	u Gln Val Gly Gly Ala A	Asn Pro Glu Val Leu Pro
1920	1925	1930
ggg ccc cgt ttc tc	c cac agc ttc ccc cgc o	gtc gga gac cac gtg gtg 5978
Gly Pro Arg Phe Se	r His Ser Phe Pro Arg \	Val Gly Asp His Val Val
. 1935	1940	1945
agc gtg cgg ggc aa	a aac cac gtg agc tgg o	gcc cag gcg cag gtg cgc 6026
Ser Val Arg Gly Ly	s Asn His Val Ser Trp A	Ala Gln Ala Gln Val Arg
1950	1955	1960
atc gtg gtg ctg ga Ile Val Val Leu Gl 1965	u Ala Val Ser Gly Leu (	cag atg ccc aac tgc tgc 6074 Gln Met Pro Asn Cys Cys 1975 1980
Glu Pro Gly Ile Al	c acg ggc act gag agg a a Thr Gly Thr Glu Arg i 85 1990	aac ttc aca gcc cgc gtg 6122 Asn Phe Thr Ala Arg Val 1995
cag cgc ggc tct cg	g gtc gcc tac gcc tgg g	tac ttc tcg ctg cag aag 6170
Gln Arg Gly Ser Ar	g Val Ala Tyr Ala Trp g	Tyr Phe Ser Leu Gln Lys
2000	2005	2010
gtc cag ggc gac to	g ctg gtc atc ctg tcg	ggc cgc gac gtc acc tac 6218
Val Gln Gly Asp Se	r Leu Val Ile Leu Ser	Gly Arg Asp Val Thr Tyr
2015	2020	2025
acg ccc gtg gcc gc	g ggg ctg ttg gag atc	cag gtg cgc gcc ttc aac 6266
Thr Pro Val Ala Al	a Gly Leu Leu Glu Ile	Gln Val Arg Ala Phe Asn
2030	2035	2040
gcc ctg ggc agt ga Ala Leu Gly Ser Gl 2045	u Asn Arg Thr Leu Val	ctg gag gtt cag gac gcc 6314 Leu Glu Val Gln Asp Ala 2055 2060
Val Gln Tyr Val Al	cc ctg cag agc ggc ccc a Leu Gln Ser Gly Pro 2070	tgc ttc acc aac cgc tcg 6362 Cys Phe Thr Asn Arg Ser 2075
gcg cag ttt gag go	cc gcc acc agc ccc agc	ccc cgg cgt gtg gcc tac 6410
Ala Gln Phe Glu Al	la Ala Thr Ser Pro Ser	Pro Arg Arg Val Ala Tyr
2080	2085	2090
cac tgg gac ttt gg	gg gat ggg tcg cca ggg	cag gac aca gat gag ccc 6458
His Trp Asp Phe G	Ly Asp Gly Ser Pro Gly	Gln Asp Thr Asp Glu Pro
2095	2100	2105
agg gcc gag cac to	cc tac ctg agg cct ggg	gac tac cgc gtg cag gtg 6506
Arg Ala Glu His So	er Tyr Leu Arg Pro Gly	Asp Tyr Arg Val Gln Val
2110	2115	2120
aac gcc tcc aac c	tg gtg agc ttc ttc gtg	gcg cag gcc acg gtg acc 6554
Asn Ala Ser Asn L	eu Val Ser Phe Phe Val	Ala Gln Ala Thr Val Thr
2125	2130	2135 2140
Val Gln Val Leu A	cc tgc cgg gag ccg gag la Cys Arg Glu Pro Glu 145 2150	gtg gac gtg gtc ctg ccc 6602 Val Asp Val Val Leu Pro 0 2155
ctg cag gtg ctg a	tg cgg cga tca cag cgc	aac tac ttg gag gcc cac 6650
Leu Gln Val Leu M	et Arg Arg Ser Gln Arg	Asn Tyr Leu Glu Ala His
2160	2165	2170
gtt gac ctg cgc g	ac tgc gtc acc tac cag	act gag tac cgc tgg gag 6698
Val Asp Leu Arg A	sp Cys Val Thr Tyr Gln	Thr Glu Tyr Arg Trp Glu

				**								_
2	2175		218	0				2185				_
gtg tat c Val Tyr A 2190		Ala Ser				Gly		Pro				6746
gcc ctg c Ala Leu P 2205			Val Ser		Pro		Leu					6794
ctg gcg c Leu Ala L				Cys		Val					Phe	6842
ggg gac a Gly Asp T	acg cca Thr Pro 2240	Leu Thr	cag agc Gln Ser	atc Ile 2245	Gln	gcc Ala	aat Asn	gtg Val	acg Thr 2250	Val	gcc Ala	6890
ccc gag c Pro Glu A 2				Glu					Arg			6938
tca gac a Ser Asp T 2270								Ser				6986
aac ctg c Asn Leu C 2285			Gln Thr				Phe					7034
gtg gct t Val Ala S						Cys					Gly	7082
ccc cgc ( Pro Arg (		Ser Thr			Pro					Ala		7130
ggc gtg g Gly Val (	gag tac Glu Tyr 2335	acc ttc Thr Phe	agc cto Ser Let 234	Thr	gtg Val	tgg Trp	aag Lys	gcc Ala 2345	Gly	cgc Arg	aag Lys	7178
gag gag g Glu Glu A 2350								Gly				7226
att gtg t Ile Val S 2365			Val Se				Gln					7274
gtg agc ( Val Ser i	cgc agc Arg Ser	tcc tac Ser Tyr 2385	gtg tac Val Ty	ttg Leu	gag Glu 239	Gly	cgc Arg	tgc Cys	ctc Leu	aat Asn 239	Cys	7322
agc agc ( Ser Ser (		Lys Arg			Ala					Ser		7370
aag acg Lys Thr	ctg gtg Leu Val 2415	ctg gat Leu Asp	gag acc Glu Th	r Thr	aca Thr	tcc Ser	acg Thr	ggc Gly 242	Ser	gca Ala	ggc Gly	7418
atg cga Met Arg 2430	Leu Val							Gly				7466

							j	LZ								
	ttc Phe 5					Leu					Glu					7514
	tcc Ser				Ser					Pro					Cys	7562
cgc Arg	ctc Leu	ttc Phe	cca Pro 2480	Leu	ggc Gly	gct Ala	gtg Val	cac His 2485	Ala	ctc Leu	acc Thr	acc Thr	aag Lys 2490	Val	cac His	7610
	gaa Glu		Thr					Ala					Ala			7658
gtg Val	tac Tyr 2510	Ala	ctg Leu	ctg Leu	ctg Leu	cgg Arg 2515	Arg	tgt Cys	cgc Arg	cag Gln	ggc Gly 252	His	tgc Cys	gag Glu	gag Glu	7706
	tgt Cys 5					Ser					Gly					7754
	ggt Gly				His					Leu					Gln	7802
	cag Gln			Ala					Leu					Ala		7850
	ctc Leu		Glu					Ala					Val			7898
cac His	ggg Gly 259	Leu	acc Thr	gct Ala	agt Ser	gtg Val 259	Leu	cca Pro	ggg Gly	ctg Leu	ctg Leu 260	Arg	cag Gln	gcc Ala	gat Asp	7946
ecc Pro 260	cag Gln	cac His	gtc Val	atc Ile	gag Glu 261	Tyr	tcg Ser	ttg Leu	gcc Ala	ctg Leu 261	Val	acc	gtg Val	ctg Leu	aac Asn 2620	7994
gaq Glu	tac Tyr	gag Glu	cgg Arg	gcc Ala 262	Leu	gac Asp	gtg Val	gcg Ala	gca Ala 263	Glu	ccc	: aaç : Lys	cac His	gag Glu 263	Arg	8042
caq Gl:	g cac n His	cga Arg	gcc Ala 264	Gln	ata Ile	cgc Arg	aag Lys	aac Asn 264	Ile	acg Thr	gaç Glu	act Thr	cto Leu 265	ı Val	tcc Ser	8090
ct ( Le	g agg 1 Arg	gtc Val 265	His	act Thr	gtg Val	gat Asp	gac Asp 266	Ile	cag Gln	cag Gln	g ato	gct Ala 266	a Ala	gcg Ala	ctg Leu	8138
gc Ala	c cag a Gln 267	Cys	ato Met	ggg Gly	ccc	ago Ser 267	Arg	gag Glu	cto Leu	gta Val	tgo Cys 268	s Ar	g Sei	g tgc Cys	ctg. Leu	8186
aa Ly: 26	g cag s Gln 35	acg	cto Lev	g cac 1 His	aag Lys 269	Leu	gaç Glu	gcc Ala	ato Met	ato Met	: Le	ato i Ile	c cto e Lei	g cag ı Glr	gca Ala 2700	8234
ga Gl	g acc u Thr	acc Thr	gcç Ala	a Gly	acc Thr	gtç Val	acq Thi	g ccc Pro	aco Thi	gco Ala	a ato	c gga e Gl	a gad y Asi	c ago o Ser	atc Ile	8282

	13		
2705	:	2710	2715
ctc aac atc aca gga g Leu Asn Ile Thr Gly A 2720			Val Arg
gca cca cag ccc tca g Ala Pro Gln Pro Ser G · 2735			
gcg tcc cag gcc tac a Ala Ser Gln Ala Tyr A 2750			
cgc tcc cgc gtg ctc a Arg Ser Arg Val Leu A 2765 2	aac gag gag ccc Asn Glu Glu Pro 2770	ctg acg ctg gcg ggc Leu Thr Leu Ala Gly 2775	gag gag 8474 Glu Glu 2780
atc gtg gcc cag ggc a Ile Val Ala Gln Gly I 2785	Lys Arg Ser Asp	ccg cgg agc ctg ctg Pro Arg Ser Leu Leu 2790	tgc tat 8522 Cys Tyr 2795
ggc ggc gcc cca ggg c Gly Gly Ala Pro Gly F 2800	cct ggc tgc cac Pro Gly Cys His 2805	ttc tcc atc ccc gag Phe Ser Ile Pro Glu 281	Ala Phe
agc ggg gcc ctg gcc a Ser Gly Ala Leu Ala A 2815			
gtg gac tcc aat ccc t Val Asp Ser Asn Pro E 2830			
tcc acc aag gtg gcc t Ser Thr Lys Val Ala S 2845			
atc ccc atc gag cgg ( Ile Pro Ile Glu Arg I 2865			
ccc aac aac tcg gac t Pro Asn Asn Ser Asp 1 2880	tgg gct gcc cgg Trp Ala Ala Arg 2885	Gly His Arg Ser Ser	Ala Asn
tcc gcc aac tcc gtt o Ser Ala Asn Ser Val v 2895			
gtc acc ctg gac agc of Val Thr Leu Asp Ser 2910	agc aac cct gcg Ser Asn Pro Ala 2915	gcc ggg ctg cat ctc Ala Gly Leu His Leu 2920	cag ctc 8906 Gln Leu
aac tat acg ctg ctg Asn Tyr Thr Leu Leu 2925			
tac ctg gca gtc tac Tyr Leu Ala Val Tyr 2945	Leu His Ser Glu	ccc cgg ccc aat gag Pro Arg Pro Asn Glu 2950	g cac aac 9002 1 His Asn 2955
tgc tcg gct agc agg Cys Ser Ala Ser Arg 2960	agg atc cgc cca Arg Ile Arg Pro 296	Glu Ser Leu Gln Gl	/ Ala Asp

							14	4						- 0	-, 0 0 , , , 20	•••
cac His	cgg Arg	ccc Pro 2975	Tyr	acc Thr	ttc Phe	ttc Phe	att Ile 2980	Ser	ccg Pro	ggg	agc Ser	aga Arg 2985	Asp	cca Pro	gcg Ala	9098
Gjy Ggg	agt Ser 2990	Tyr	cat His	ctg Leu	aac Asn	ctc Leu 2995	Ser	agc Ser	cac His	ttc Phe	cgc Arg 3000	Trp	tcg Ser	gcg Ala	ctg Leu	9146
cag Gln 3005	Val	tcc Ser	gtg Val	ggc Gly	ctg Leu 3010	Tyr	acg Thr	tcc Ser	ctg Leu	tgc Cys 3015	cag Gln	tac Tyr	ttc Phe	agc Ser	gag Glu 3020	9194
gag Glu	gac Asp	atg Met	gtg Val	tgg Trp 3025	Arg	aca Thr	gag Glu	GJ À aàà	ctg Leu 3030	Leu	ccc Pro	ctg Leu	gag Glu	gag Glu 3035	Thr	9242
tcg Ser	ccc Pro	cgc Arg	cag Gln 3040	Ala	gtc Val	tgc Cys	ctc Leu	acc Thr 3045	Arg	cac His	ctc Leu	acc Thr	gcc Ala 3050	Phe	ggc Gly	9290
gcc Ala	agc Ser	ctc Leu 3055	Phe	gtg Val	ccc Pro	cca Pro	agc Ser 3060	His	gtc Val	cgc Arg	ttt Phe	gtg Val 3065	Phe	cct Pro	gag Glu	9338
ccg Pro	aca Thr 3070	Ala	gat Asp	gta Val	aac Asn	tac Tyr 3075	Ile	gtc Val	atg Met	ctg Leu	aca Thr 3080	Cys	gct Ala	gtg Val	tgc Cys	9386
ctg Leu 3085	Val	acc Thr	tac Tyr	atg Met	gtc Val 3090	Met	gcc Ala	gcc Ala	atc Ile	ctg Leu 3095	cac His	aag Lys	ctg Leu	gac Asp	cag Gln 3100	9434
ttg Leu	gat Asp	gcc Ala	agc Ser	cgg Arg 3109	Gly	cgc Arg	gcc Ala	atc Ile	cct Pro 3110	Phe	tgt Cys	ggg Gly	cag Gln	cgg Arg 3115	Gly	9482
cgc Arg	ttc Phe	aag Lys	tac Tyr 3120	Glu	atc Ile	ctc Leu	gtc Val	aag Lys 3125	Thr	ggc Gly	tgg Trp	ggc Gly	cgg Arg 3130	Gly	tca Ser	9530
ggt Gly	acc Thr	acg Thr 3135	Ala	cac His	gtg Val	ggc Gly	atc Ile 3140	Met	ctg Leu	tat Tyr	Gly ggg	gtg Val 3145	Asp	agc Ser	cgg Arg	9578
agc Ser	ggc Gly 3150	His	cgg Arg	cac His	ctg Leu	gac Asp 3155	Gly	gac Asp	aga Arg	gcc Ala	ttc Phe 3160	His	cgc Arg	aac Asn	agc Ser	9626
ctg Leu 3165	Asp	atc Ile	ttc Phe	cgg Arg	atc Ile 3170	Ala	acc Thr	ccg Pro	cac His	agc Ser 317	ctg Leu 5	ggt Gly	agc Ser	gtg Val	tgg Trp 3180	9674
aag Lys	atc Ile	cga Arg	gtg Val	tgg Trp 318	His	gac Asp	aac Asn	aaa Lys	ggg Gly 319	Leu	agc Ser	cct Pro	gcc Ala	tgg Trp 319	Phe	9722
ctg Leu	cag Gln	cac His	gtc Val 3200	Ile	gtc Val	agg Arg	gac Asp	ctg Leu 320	Gln	acg Thr	gca Ala	cgc Arg	agc Ser 321	Ala	ttc Phe	9770
ttc Phe	ctg Leu	gtc Val 321	Asn	gac Asp	tgg Trp	ctt Leu	tcg Ser 3220	Val	gag Glu	acg Thr	gag Glu	gcc Ala 322	Asn	ggg Gly	ggc Gly	9818
ctg Leu	gtg Val	gag Glu	aag Lys	gag Glu	gtg Val	ctg Leu	gcc Ala	gcg Ala	agc Ser	gac Asp	gca Ala	gcc Ala	ctt Leu	ttg Leu	cgc Arg	9866

								13								
	3230					3235					3240	)				-
ttc Phe 3245	Arg	cgc Arg	ctg Leu	ctg Leu	gtg Val 3250	Ala	gag Glu	ctg Leu	cag Gln	cgt Arg 3255	Gly	ttc Phe	ttt Phe	gac Asp	aag Lys 3260	9914
cac His	atc Ile	tgg Trp	ctc Leu	tcc Ser 3265	Ile	tgg Trp	gac Asp	cgg Arg	ccg Pro 3270	Pro	cgt Arg	agc Ser	cgt Arg	ttc Phe 32 <b>7</b> 5	Thr	9962
cgc Arg	atc Ile	cag Gln	agg Arg 3280	Ala	acc Thr	tgc Cys	tgc Cys	gtt Val 3285	Leu	ctc Leu	atc Ile	tgc Cys	ctc Leu 3290	Phe	ctg Leu	10010
ggc Gly	gcc Ala	aac Asn 3295	Ala	gtg Val	tgg Trp	tac Tyr	ggg Gly 3300	Ala	gtt Val	ggc Gly	gac Asp	tct Ser 3305	Ala	tac Tyr	agc Ser	10058
acg Thr	ggg Gly 3310	His	gtg Val	tcc Ser	agg Arg	ctg Leu 331	Ser	ccg Pro	ctg Leu	agc Ser	gtc Val 332	gac Asp 0	aca Thr	gtc Val	gct Ala	10106
gtt Val 332	Gly	ctg Leu	gtg Val	tcc Ser	agc Ser 3330	Val	gtt Val	gtc Val	tat Tyr	ccc Pro 333	Val	tac Tyr	ctg Leu	gcc Ala	atc Ile 3340	10154
ctt Leu	ttt Phe	ctc Leu	ttc Phe	cgg Arg 334	Met	tcc Ser	cgg Arg	agc Ser	aag Lys 335	Val	gct Ala	ggģ Gly	agc Ser	ccg Pro 335	Ser	10202
ccc Pro	aca Thr	cct Pro	gcc Ala 336	Gly	cag Gln	cag Gln	gtg Val	ctg Leu 336	Asp	atc Ile	gac Asp	agc Ser	tgc Cys 337	Leu	gac Asp	10250
tcg Ser	tcc Ser	gtg Val 337	Leu	gac Asp	agc Ser	tcc Ser	ttc Phe 338	Leu	acg Thr	ttc Phe	tca Ser	ggc Gly 338	Leu	cac His	gct Ala	10298
gag Glu	gcc Ala 339	Phe	gtt Val	gga Gly	cag Gln	atg Met 339	Lys	agt Ser	gac Asp	tto Lev	ttt Phe 340	ctg Leu	gat Asp	gat Asp	tct Ser	10346
aag Lys 340	Ser	ctg Leu	gtg Val	tgc Cys	tgg Trp 341	Pro	tcc Ser	: ggc	gag Glu	g gga Gly 341	Thr	g ctc Leu	agt Ser	tgg Trp	ccg Pro 3420	10394
gac Asp	ctg Leu	cto Leu	: agt : Ser	gac Asp 342	) Pro	tcc Ser	: att	gto Val	g ggt L Gly 343	, Sei	c aat r Ası	ctg n Leu	Arg	g caç g Glr 343	ı Leu	10442
gca Ala	a cgg a Arg	GJ?	caç Glr 344	ı Ala	g ggc Gly	cat His	: ggg	g cto / Let 344	ı Gly	c cca y Pro	a gaq o Gli	g gag u Glu	gac Asp 345	GT?	ttc y Phe	10490
tc: Se:	c ctq r Leu	g gcc Ala 345	a Ser	c ccc	tac Tyr	tco Sei	g cct Pro 340	o Ala	c aaa a Lys	a tc s Se	c tte r Ph	c tca e Ser 346	Ala	a tca a Sei	a gat r Asp	10538
gaa Gl:	a gad u Asp 347	) Le	g ato ı Ile	c caq e Glr	g caq n Glr	g gto Vai 34	L Le	t gc	c gae a Gl	g gg u Gl	g gt y Va 34	l Ser	ago Se:	c cca	a gcc o Ala	10586
cc Pr	o Thi	c caa	a gad n Ası	c acc	c cad r His 349	s Me	g ga t Gl	a ac u Th	g ga r As	p Le	g ct u Le 95	c ago u Sei	c age	c ct r Le	g tcc u Ser 3500	10634

	ggg gag aag aca Gly Glu Lys Thr 3505			Gly
Glu Leu Gly	cca ccc agc cca Pro Pro Ser Pro 3520			
	tcc agg aca gga Ser Arg Thr Gly			
ctg ccg gcc Leu Pro Ala 3550	tgg tgt gcc tcc Trp Cys Ala Ser 3555	Leu Ala His Gly	ctc agc ctg ctc Leu Ser Leu Leu 3560	ctg 10826 Leu
	gct gtg gct gtc Ala Val Ala Val 3570		Gly Ala Ser Phe	
	agt gtt gcg tgg Ser Val Ala Trp 3585			Leu
	ctc ggc tgg gag Leu Gly Trp Glu 3600			
tac ttc tca Tyr Phe Ser 3615	ctg gtg gcc aag Leu Val Ala Lys	cgg ctg cac ccg Arg Leu His Pro 3620	gat gaa gat gac Asp Glu Asp Asp 3625	acc 11018 Thr
	agc ccg gct gtg Ser Pro Ala Val 3635	Thr Pro Val Ser		
	ccc cac ggc ttt Pro His Gly Phe 3650		Ala Lys Glu Glu	
	aag agg cta cat Lys Arg Leu His 3665			Tyr
atg ctt ttt Met Leu Phe	ctg ctg gtg acc Leu Leu Val Thr 3680	ctg ctg gcc agc Leu Leu Ala Ser 3685	tat ggg gat gcc Tyr Gly Asp Ala 3690	tca 11210 Ser
tgc cat ggg Cys His Gly 3695	cac gcc tac cgt His Ala Tyr Arg 5	ctg caa agc gcc Leu Gln Ser Ala 3700	atc aag cag gag Ile Lys Gln Glu 3705	ctg 11258 Leu
cac agc cgg His Ser Arg 3710	gcc ttc ctg gcc Ala Phe Leu Ala 371	Ile Thr Arg Ser	gag gag ctc tgg Glu Glu Leu Trp 3720	cca 11306 Pro
	cac gtg ctg ctg His Val Leu Leu 3730		Gly Asn Gln Ser	
cca gag ctg Pro Glu Leu	ggg ccc cca cgg Gly Pro Pro Arg 3745	ctg cgg cag gtg Leu Arg Gln Val 3750	cgg ctg cag gaa Arg Leu Gln Glu 379	ı Ala
ctc tac cca Leu Tyr Pro	gac cct ccc ggc Asp Pro Pro Gly	ccc agg gtc cac Pro Arg Val His	e acg tgc tcg gcc s Thr Cys Ser Ala	c gca 11450 a Ala

	17	
3760	3765	3770
gga ggc ttc agc acc	agc gat tac gac gtt ggc tgg ga	g agt cct cac 11498
Gly Gly Phe Ser Thr	Ser Asp Tyr Asp Val Gly Trp Gl	u Ser Pro His
3775	3780	'85
aat ggc tcg ggg acg Asn Gly Ser Gly Thr 3790	tgg gcc tat tca gcg ccg gat ct Trp Ala Tyr Ser Ala Pro Asp Le 3795 3800	g ctg ggg gca 11546 eu Leu Gly Ala
tgg tcc tgg ggc tcc	tgt gcc gtg tat gac agc ggg gg	gc tac gtg cag 11594
Trp Ser Trp Gly Ser	Cys Ala Val Tyr Asp Ser Gly Gl	Ly Tyr Val Gln
3805	3810 3815	3820
gag ctg ggc ctg agc	ctg gag gag agc cgc gac cgg ct	eg cgc ttc ctg 11642
Glu Leu Gly Leu Ser	Leu Glu Glu Ser Arg Asp Arg Le	eu Arg Phe Leu
3825	3830	3835
cag ctg cac aac tgg	ctg gac aac agg agc cgc gct gt	tg ttc ctg gag 11690
Gln Leu His Asn Trp	Leu Asp Asn Arg Ser Arg Ala Va	al Phe Leu Glu
3840	3845	3850
ctc acg cgc tac agc	ccg gcc gtg ggg ctg cac gcc g	cc gtc acg ctg 11738
Leu Thr Arg Tyr Ser	Pro Ala Val Gly Leu His Ala A	la Val Thr Leu
3855	3860	865
cgc ctc gag ttc ccg Arg Leu Glu Phe Pro 3870	gcg gcc ggc cgc gcc ctg gcc g Ala Ala Gly Arg Ala Leu Ala A 3875 3880	cc ctc agc gtc 11786 la Leu Ser Val
cgc ccc ttt gcg ctg	cgc cgc ctc agc gcg ggc ctc t	cg ctg cct ctg 11834
Arg Pro Phe Ala Leu	Arg Arg Leu Ser Ala Gly Leu S	er Leu Pro Leu
3885	3890 3895	3900
ctc acc tcg gtg tgc	ctg ctg ctg ttc gcc gtg cac t	tc gcc gtg gcc 11882
Leu Thr Ser Val Cys	Leu Leu Leu Phe Ala Val His P	he Ala Val Ala
390	5 3910	3915
gag gcc cgt act tgg	cac agg gaa ggg cgc tgg cgc g	tg ctg cgg ctc 11930
Glu Ala Arg Thr Trp	His Arg Glu Gly Arg Trp Arg V	'al Leu Arg Leu
3920	3925	3930
gga gcc tgg gcg cgg	tgg ctg ctg gtg gcg ctg acg g	geg gee aeg gea 11978
Gly Ala Trp Ala Arg	Trp Leu Leu Val Ala Leu Thr A	Ala Ala Thr Ala
3935	3940	8945
ctg gta cgc ctc gcc Leu Val Arg Leu Ala 3950	cag ctg ggt gcc gct gac cgc o Gln Leu Gly Ala Ala Asp Arg 0 3955 3960	cag tgg acc cgt 12026 Gln Trp Thr Arg
ttc gtg cgc ggc cgc	ccg cgc cgc ttc act agc ttc q	gac cag gtg gcg 12074
Phe Val Arg Gly Arg	g Pro Arg Arg Phe Thr Ser Phe A	Asp Gln Val Ala
3965	3970 3975	3980
cag ctg agc tcc gca	a gcc cgt ggc ctg gcg gcc tcg o	ctg ctc ttc ctg 12122
Gln Leu Ser Ser Ala	a Ala Arg Gly Leu Ala Ala Ser 1	Leu Leu Phe Leu
398	35 3990	3995
ctt ttg gtc aag gct	gcc cag cag cta cgc ttc gtg o	age cag tgg tee 12170
Leu Leu Val Lys Ala	Ala Gln Gln Leu Arg Phe Val 2	Arg Gln Trp Ser
4000	4005	4010
gtc ttt ggc aag aca	a tta tgc cga gct ctg cca gag	ctc ctg ggg gtc 12218
Val Phe Gly Lys Thr	r Leu Cys Arg Ala Leu Pro Glu	Leu Leu Gly Val
4015	4020	4025

							]	18						• `	21100771	25071
		Gly				ctc Leu 4035	Gly					Gln				12266
	Leu					gtg Val					Ser					12314
					Pro	ggg Gly				Ser					Ala	12362
				Leu		ccc Pro			Cys					Āla		12410
			Gly			cgg Arg		Gly					Arg			12458
		Āla				gag Glu 4115	Leu					Trp				12506
	Tyr					ttg Leu )					Leu					12554
					Lys	gag Glu				Lys					Gly	12602
				Pro		cgc Arg			Arg					Ser		12650
			Pro			gct Ala		Ser					Pro			12698
		Ser				ggg Gly 4195	Leu					Gly				12746
	Arg					ccc Pro 0					Āla					12794
					Asp	cga Arg				Āla					Tyr	12842
				Gln		cac His			Gln					Ser		12890
			Gly			cgt Arg		Pro					Arg			12938
		Ser				cgg Arg 427	Ala					Asp				12986
ggc	ccc Pro	agc Ser	agg Arg	aca Thr	ccc Pro	ctt Leu	ċgg Arg	gcc Ala	aag Lys	aac Asn	aag Lys	gtc Val	cac His	ccc Pro	agc Ser	13034

4285 4295 4290 4300 age act tagteeteet teetggeggg ggtgggeegt ggagteggag tggacaeege 13090 Ser Thr tragtattac tttctgccgc tgtcaaggcc gagggccagg cagaatggct gcacgtaggt 13150 tececagaga geaggeaggg geatetgtet gtetgtggge tteageactt taaagagget 13210 gtgtggccaa ccaggaccca gggtcccctc cccagctccc ttgggaagga cacagcagta 13270 ttggacggtt tctagcctct gagatgctaa tttatttccc cgagtcctca ggtacagcgg 13330 gctgtgcccg gccccaccc ctgggcagat gtcccccact gctaaggctg ctggcttcag 13390 ggagggttag cctgcaccgc cgccaccctg cccctaagtt attacctctc cagitcctac 13450 cgtactccct gcaccgtctc actgtgtgtc tcgtgtcagt aatttatatg gtgttaaaat gigtatattt tigtatgica ciatticac tagggetgag gggeetgege ecagagetgg cctcccccaa cacctgctgc gcttggtagg tgtggtggcg ttatggcagc ccggctgctg cttggatgcg agcttggcct tgggccggtg ctgggggcac agctgtctgc caggcactct 13690 catcacccca gaggeettgt catceteeet tgeeccagge caggtageaa gagageageg 13750 cccaggcctg ctggcatcag gtctgggcaa gtagcaggac taggcatgtc agaggacccc 13810 agggtggtta gaggaaaaga ctcctcctgg gggctggctc ccagggtgga ggaaggtgac 13870 tgtgtgtgtg tgtgtgtgcg cgcgcgcacg cgcgagtgtg ctgtatggcc caggcagcct caaggeeete ggagetgget gtgeetgett etgtgtacea ettetgtggg eatggeeget totagageet egacaceee ecaaceeeg caccaageag acaaagteaa taaaagaget 14050 gtctgactgc 14060 <210> 2 <211> 4302 <212> PRT <213> Homo sapiens <400> 2 Met Pro Pro Ala Ala Pro Ala Arg Leu Ala Leu Ala Leu Gly Leu Gly 10 Leu Trp Leu Gly Ala Leu Ala Gly Gly Pro Gly Arg Gly Cys Gly Pro 20 25 Cys Glu Pro Pro Cys Leu Cys Gly Pro Ala Pro Gly Ala Ala Cys Arg 40 Val Asn Cys Ser Gly Arg Gly Leu Arg Thr Leu Gly Pro Ala Leu Arg 55 Ile Pro Ala Asp Ala Thr Ala Leu Asp Val Ser His Asn Leu Leu Arg 70 7.5 Ala Leu Asp Val Gly Leu Leu Ala Asn Leu Ser Ala Leu Ala Glu Leu Asp Ile Ser Asn Asn Lys Ile Ser Thr Leu Glu Glu Gly Ile Phe Ala 100 105 Asn Leu Phe Asn Leu Ser Glu Ile Asn Leu Ser Gly Asn Pro Phe Glu 120 125 Cys Asp Cys Gly Leu Ala Trp Leu Pro Arg Trp Ala Glu Gln Gln 130 135 140 Val Arg Val Val Gln Pro Glu Ala Ala Thr Cys Ala Gly Pro Gly Ser 150 155 160 Leu Ala Gly Gln Pro Leu Leu Gly Ile Pro Leu Leu Asp Ser Gly Cys 165 170 175 Gly Glu Glu Tyr Val Ala Cys Leu Pro Asp Asn Ser Ser Gly Thr Val 180 185 190 Ala Ala Val Ser Phe Ser Ala Ala His Glu Gly Leu Leu Gln Pro Glu 195 200 205 Ala Cys Ser Ala Phe Cys Phe Ser Thr Gly Gln Gly Leu Ala Ala Leu 220 Ser Glu Gln Gly Trp Cys Leu Cys Gly Ala Ala Gln Pro Ser Ser Ala 230 235 Ser Phe Ala Cys Leu Ser Leu Cys Ser Gly Pro Pro Pro Pro Ala 245 250 Pro Thr Cys Arg Gly Pro Thr Leu Leu Gln His Val Phe Pro Ala Ser 265 Pro Gly Ala Thr Leu Val Gly Pro His Gly Pro Leu Ala Ser Gly Gln

280

285

275

Leu Ala Ala Phe His Ile Ala Ala Pro Leu Pro Val Thr Ala Thr Arg Trp Asp Phe Gly Asp Gly Ser Ala Glu Val Asp Ala Ala Gly Pro Ala Ala Ser His Arg Tyr Val Leu Pro Gly Arg Tyr His Val Thr Ala Val Leu Ala Leu Gly Ala Gly Ser Ala Leu Leu Gly Thr Asp Val Gln Val Glu Ala Ala Pro Ala Ala Leu Glu Leu Val Cys Pro Ser Ser Val Gln Ser Asp Glu Ser Leu Asp Leu Ser Ile Gln Asn Arg Gly Gly Ser Gly Leu Glu Ala Ala Tyr Ser Ile Val Ala Leu Gly Glu Glu Pro Ala Arg Ala Val His Pro Leu Cys Pro Ser Asp Thr Glu Ile Phe Pro Gly Asn Gly His Cys Tyr Arg Leu Val Val Glu Lys Ala Ala Trp Leu Gln Ala Gln Glu Gln Cys Gln Ala Trp Ala Gly Ala Ala Leu Ala Met Val Asp Ser Pro Ala Val Gln Arg Phe Leu Val Ser Arg Val Thr Arg Ser Leu Asp Val Trp Ile Gly Phe Ser Thr Val Gln Gly Val Glu Val Gly Pro Ala Pro Gln Gly Glu Ala Phe Ser Leu Glu Ser Cys Gln Asn Trp Leu Pro Gly Glu Pro His Pro Ala Thr Ala Glu His Cys Val Arg Leu Gly Pro Thr Gly Trp Cys Asn Thr Asp Leu Cys Ser Ala Pro His Ser Tyr Val Cys Glu Leu Gln Pro Gly Gly Pro Val Gln Asp Ala Glu Asn Leu Leu Val Gly Ala Pro Ser Gly Asp Leu Gln Gly Pro Leu Thr Pro Leu Ala Gln Gln Asp Gly Leu Ser Ala Pro His Glu Pro Val Glu Val Met Val Phe Pro Gly Leu Arg Leu Ser Arg Glu Ala Phe Leu Thr Thr Ala Glu Phe Gly Thr Gln Glu Leu Arg Arg Pro Ala Gln Leu Arg Leu Gln Val Tyr Arg Leu Leu Ser Thr Ala Gly Thr Pro Glu Asn Gly Ser Glu Pro Glu Ser Arg Ser Pro Asp Asn Arg Thr Gln Leu Ala Pro Ala Cys Met Pro Gly Gly Arg Trp Cys Pro Gly Ala Asn Ile Cys Leu Pro Leu Asp Ala Ser Cys His Pro Gln Ala Cys Ala Asn Gly Cys Thr Ser Gly Pro Gly Leu Pro Gly Ala Pro Tyr Ala Leu Trp Arg Glu Phe Leu Phe Ser Val Ala Ala Gly Pro Pro Ala Gln Tyr Ser Val Thr Leu His Gly Gln Asp Val Leu Met Leu Pro Gly Asp Leu Val Gly Leu Gln His Asp Ala Gly Pro Gly Ala Leu Leu His Cys Ser Pro Ala Pro Gly His Pro Gly Pro Gln Ala Pro Tyr Leu Ser Ala Asn Ala Ser Ser Trp Leu Pro His Leu Pro Ala Gln Leu Glu Gly Thr Trp Ala Cys Pro Ala Cys Ala Leu Arg Leu Leu Ala Ala Thr Glu Gln Leu Thr Val Leu Leu Gly Leu Arg Pro Asn Pro Gly Leu Arg Met Pro Gly Arg Tyr Glu Val Arg Ala Glu Val Gly Asn Gly Val Ser Arg His Asn Leu Ser Cys Ser Phe Asp 

Val Val Ser Pro Val Ala Gly Leu Arg Val Ile Tyr Pro Ala Pro Arg Asp Gly Arg Leu Tyr Val Pro Thr Asn Gly Ser Ala Leu Val Leu Gln Val Asp Ser Gly Ala Asn Ala Thr Ala Thr Ala Arg Trp Pro Gly Gly Ser Val Ser Ala Arg Phe Glu Asn Val Cys Pro Ala Leu Val Ala Thr Phe Val Pro Gly Cys Pro Trp Glu Thr Asn Asp Thr Leu Phe Ser Val Val Ala Leu Pro Trp Leu Ser Glu Gly Glu His Val Val Asp Val Val Val Glu Asn Ser Ala Ser Arg Ala Asn Leu Ser Leu Arg Val Thr Ala Glu Glu Pro Ile Cys Gly Leu Arg Ala Thr Pro Ser Pro Glu Ala Arg Val Leu Gln Gly Val Leu Val Arg Tyr Ser Pro Val Val Glu Ala Gly Ser Asp Met Val Phe Arg Trp Thr Ile Asn Asp Lys Gln Ser Leu Thr Phe Gln Asn Val Val Phe Asn Val Ile Tyr Gln Ser Ala Ala Val Phe Lys Leu Ser Leu Thr Ala Ser Asn His Val Ser Asn Val Thr Val Asn Tyr Asn Val Thr Val Glu Arg Met Asn Arg Met Gln Gly Leu Gln Val 1010 1015 1020 Ser Thr Val Pro Ala Val Leu Ser Pro Asn Ala Thr Leu Ala Leu Thr Ala Gly Val Leu Val Asp Ser Ala Val Glu Val Ala Phe Leu Trp Asn 1050 1055 Phe Gly Asp Gly Glu Gln Ala Leu His Gln Phe Gln Pro Pro Tyr Asn · 1065 1070 Glu Ser Phe Pro Val Pro Asp Pro Ser Val Ala Gln Val Leu Val Glu 1080 1085 His Asn Val Met His Thr Tyr Ala Ala Pro Gly Glu Tyr Leu Leu Thr Val Leu Ala Ser Asn Ala Phe Glu Asn Leu Thr Gln Gln Val Pro Val Ser Val Arg Ala Ser Leu Pro Ser Val Ala Val Gly Val Ser Asp Gly Val Leu Val Ala Gly Arg Pro Val Thr Phe Tyr Pro His Pro Leu Pro Ser Pro Gly Gly Val Leu Tyr Thr Trp Asp Phe Gly Asp Gly Ser Pro Val Leu Thr Gln Ser Gln Pro Ala Ala Asn His Thr Tyr Ala Ser Arg 1170 1175 1180 Gly Thr Tyr His Val Arg Leu Glu Val Asn Asn Thr Val Ser Gly Ala Ala Ala Gln Ala Asp Val Arg Val Phe Glu Glu Leu Arg Gly Leu Ser Val Asp Met Ser Leu Ala Val Glu Gln Gly Ala Pro Val Val Ser Ala Ala Val Gln Thr Gly Asp Asn Ile Thr Trp Thr Phe Asp Met Gly Asp Gly Thr Val Leu Ser Gly Pro Glu Ala Thr Val Glu His Val Tyr 1250 1255 Leu Arg Ala Gln Asn Cys Thr Val Thr Val Gly Ala Ala Ser Pro Ala Gly His Leu Ala Arg Ser Leu His Val Leu Val Phe Val Leu Glu Val Leu Arg Val Glu Pro Ala Ala Cys Ile Pro Thr Gln Pro Asp Ala Arg Leu Thr Ala Tyr Val Thr Gly Asn Pro Ala His Tyr Leu Phe Asp Trp Thr Phe Gly Asp Gly Ser Ser Asn Thr Thr Val Arg Gly Cys Pro Thr 

Val Thr His Asn Phe Thr Arg Ser Gly Thr Phe Pro Leu Ala Leu Val 1355 1360 Leu Ser Ser Arg Val Asn Arg Ala His Tyr Phe Thr Ser Ile Cys Val Glu Pro Glu Val Gly Asn Val Thr Leu Gln Pro Glu Arg Gln Phe Val 1385 1390 Gln Leu Gly Asp Glu Ala Trp Leu Val Ala Cys Ala Trp Pro Pro Phe Pro Tyr Arg Tyr Thr Trp Asp Phe Gly Thr Glu Glu Ala Ala Pro Thr Arg Ala Arg Gly Pro Glu Val Thr Phe Ile Tyr Arg Asp Pro Gly Ser Tyr Leu Val Thr Val Thr Ala Ser Asn Asn Ile Ser Ala Ala Asn Asp Ser Ala Leu Val Glu Val Gln Glu Pro Val Leu Val Thr Ser Ile Lys Val Asn Gly Ser Leu Gly Leu Glu Leu Gln Gln Pro Tyr Leu Phe Ser 1475 1480 1485 Ala Val Gly Arg Gly Arg Pro Ala Ser Tyr Leu Trp Asp Leu Gly Asp 1495 1500 Gly Gly Trp Leu Glu Gly Pro Glu Val Thr His Ala Tyr Asn Ser Thr Gly Asp Phe Thr Val Arg Val Ala Gly Trp Asn Glu Val Ser Arg Ser Glu Ala Trp Leu Asn Val Thr Val Lys Arg Arg Val Arg Gly Leu Val 1540 1545 1550 Val Asn Ala Ser Arg Thr Val Val Pro Leu Asn Gly Ser Val Ser Phe Ser Thr Ser Leu Glu Ala Gly Ser Asp Val Arg Tyr Ser Trp Val Leu 1570 1575 1580 Cys Asp Arg Cys Thr Pro Ile Pro Gly Gly Pro Thr Ile Ser Tyr Thr 1590 1595 Phe Arg Ser Val Gly Thr Phe Asn Ile Ile Val Thr Ala Glu Asn Glu 1605 1610 1615 Val Gly Ser Ala Gln Asp Ser Ile Phe Val Tyr Val Leu Gln Leu Ile Glu Gly Leu Gln Val Val Gly Gly Gly Arg Tyr Phe Pro Thr Asn His Thr Val Gln Leu Gln Ala Val Val Arg Asp Gly Thr Asn Val Ser Tyr Ser Trp Thr Ala Trp Arg Asp Arg Gly Pro Ala Leu Ala Gly Ser Gly Lys Gly Phe Ser Leu Thr Val Leu Glu Ala Gly Thr Tyr His Val Gln Leu Arg Ala Thr Asn Met Leu Gly Ser Ala Trp Ala Asp Cys Thr Met Asp Phe Val Glu Pro Val Gly Trp Leu Met Val Thr Ala Ser Pro Asn Pro Ala Ala Val Asn Thr Ser Val Thr Leu Ser Ala Glu Leu Ala Gly 1730 1735 Gly Ser Gly Val Val Tyr Thr Trp Ser Leu Glu Gly Leu Ser Trp Glu Thr Ser Glu Pro Phe Thr Thr His Ser Phe Pro Thr Pro Gly Leu His Leu Val Thr Met Thr Ala Gly Asn Pro Leu Gly Ser Ala Asn Ala Thr Val Glu Val Asp Val Gln Val Pro Val Ser Gly Leu Ser Ile Arg Ala Ser Glu Pro Gly Gly Ser Phe Val Ala Ala Gly Ser Ser Val Pro Phe Trp Gly Gln Leu Ala Thr Gly Thr Asn Val Ser Trp Cys Trp Ala Val Pro Gly Gly Ser Ser Lys Arg Gly Pro His Val Thr Met Val Phe Pro Asp Ala Gly Thr Phe Ser Ile Arg Leu Asn Ala Ser Asn Ala Val 

Ser Trp Val Ser Ala Thr Tyr Asn Leu Thr Ala Glu Glu Pro Ile Val 1875 1880 1885 Gly Leu Val Leu Trp Ala Ser Ser Lys Val Val Ala Pro Gly Gln Leu 1895 1900 Val His Phe Gln Ile Leu Leu Ala Ala Gly Ser Ala Val Thr Phe Arg 1915 1910 Leu Gln Val Gly Gly Ala Asn Pro Glu Val Leu Pro Gly Pro Arg Phe 1930 1925 1935 Ser His Ser Phe Pro Arg Val Gly Asp His Val Val Ser Val Arg Gly 1940 1945 1950 Lys Asn His Val Ser Trp Ala Gln Ala Gln Val Arg Ile Val Val Leu 1955 1960 1965 Glu Ala Val Ser Gly Leu Gln Met Pro Asn Cys Cys Glu Pro Gly Ile 1975 1980 Ala Thr Gly Thr Glu Arg Asn Phe Thr Ala Arg Val Gln Arg Gly Ser 1990 1995 Arg Val Ala Tyr Ala Trp Tyr Phe Ser Leu Gln Lys Val Gln Gly Asp 2010 2005 Ser Leu Val Ile Leu Ser Gly Arg Asp Val Thr Tyr Thr Pro Val Ala 2025 2020 2030 Ala Gly Leu Leu Glu Ile Gln Val Arg Ala Phe Asn Ala Leu Gly Ser 2040 2045 2035 Glu Asn Arg Thr Leu Val Leu Glu Val Gln Asp Ala Val Gln Tyr Val 2055 2060 Ala Leu Gln Ser Gly Pro Cys Phe Thr Asn Arg Ser Ala Gln Phe Glu 2070 2075 Ala Ala Thr Ser Pro Ser Pro Arg Arg Val Ala Tyr His Trp Asp Phe 2085 2090 2095 Gly Asp Gly Ser Pro Gly Gln Asp Thr Asp Glu Pro Arg Ala Glu His 2105 2110 2100 Ser Tyr Leu Arg Pro Gly Asp Tyr Arg Val Gln Val Asn Ala Ser Asn 2115 2120 2125 Leu Val Ser Phe Phe Val Ala Gln Ala Thr Val Thr Val Gln Val Leu 2135 2140 Ala Cys Arg Glu Pro Glu Val Asp Val Leu Pro Leu Gln Val Leu 2150 2155 Met Arg Arg Ser Gln Arg Asn Tyr Leu Glu Ala His Val Asp Leu Arg 2170 2175 2165 Asp Cys Val Thr Tyr Gln Thr Glu Tyr Arg Trp Glu Val Tyr Arg Thr 2180 2185 2190 Ala Ser Cys Gln Arg Pro Gly Arg Pro Ala Arg Val Ala Leu Pro Gly 2195 2200 2205 Val Asp Val Ser Arg Pro Arg Leu Val Leu Pro Arg Leu Ala Leu Pro 2220 2215 Val Gly His Tyr Cys Phe Val Phe Val Val Ser Phe Gly Asp Thr Pro 2230 2235 Leu Thr Gln Ser Ile Gln Ala Asn Val Thr Val Ala Pro Glu Arg Leu 2250 2255 2245 Val Pro Ile Ile Glu Gly Gly Ser Tyr Arg Val Trp Ser Asp Thr Arg 2270 2260 2265 -Asp Leu Val Leu Asp Gly Ser Glu Ser Tyr Asp Pro Asn Leu Glu Asp 2285 2275 2280 Gly Asp Gln Thr Pro Leu Ser Phe His Trp Ala Cys Val Ala Ser Thr 2300 2295 Gln Arg Glu Ala Gly Gly Cys Ala Leu Asn Phe Gly Pro Arg Gly Ser 2315 2310 Ser Thr Val Thr Ile Pro Arg Glu Arg Leu Ala Ala Gly Val Glu Tyr 2330 2325 Thr Phe Ser Leu Thr Val Trp Lys Ala Gly Arg Lys Glu Glu Ala Thr 2350 2340 2345 Asn Gln Thr Val Leu Ile Arg Ser Gly Arg Val Pro Ile Val Ser Leu 2355 2365 2360 Glu Cys Val Ser Cys Lys Ala Gln Ala Val Tyr Glu Val Ser Arg Ser 2375 2380 Ser Tyr Val Tyr Leu Glu Gly Arg Cys Leu Asn Cys Ser Ser Gly Ser 2390 2395

Lys Arg Gly Arg Trp Ala Ala Arg Thr Phe Ser Asn Lys Thr Leu Val 2405 2410 Leu Asp Glu Thr Thr Ser Thr Gly Ser Ala Gly Met Arg Leu Val 2420 2425 Leu Arg Arg Gly Val Leu Arg Asp Gly Glu Gly Tyr Thr Phe Thr Leu 2435 2440 2445 Thr Val Leu Gly Arg Ser Gly Glu Glu Glu Gly Cys Ala Ser Ile Arg 2455 2460 Leu Ser Pro Asn Arg Pro Pro Leu Gly Gly Ser Cys Arg Leu Phe Pro 2470 2475 Leu Gly Ala Val His Ala Leu Thr Thr Lys Val His Phe Glu Cys Thr 2485 2490 Gly Trp His Asp Ala Glu Asp Ala Gly Ala Pro Leu Val Tyr Ala Leu 2500 2505 2510 Leu Leu Arg Arg Cys Arg Gln Gly His Cys Glu Glu Phe Cys Val Tyr 2520 2515 2525 Lys Gly Ser Leu Ser Ser Tyr Gly Ala Val Leu Pro Pro Gly Phe Arg 2535 2540 Pro His Phe Glu Val Gly Leu Ala Val Val Gln Asp Gln Leu Gly 2550 2555 Ala Ala Val Val Ala Leu Asn Arg Ser Leu Ala Ile Thr Leu Pro Glu 2565 2570 Pro Asn Gly Ser Ala Thr Gly Leu Thr Val Trp Leu His Gly Leu Thr 2580 2585 2590 Ala Ser Val Leu Pro Gly Leu Leu Arg Gln Ala Asp Pro Gln His Val 2605 2595 2600 Ile Glu Tyr Ser Leu Ala Leu Val Thr Val Leu Asn Glu Tyr Glu Arg 2615 2620 Ala Leu Asp Val Ala Ala Glu Pro Lys His Glu Arg Gln His Arg Ala 2630 2635 Gln Ile Arg Lys Asn Ile Thr Glu Thr Leu Val Ser Leu Arg Val His 2645 2650 Thr Val Asp Asp Ile Gln Gln Ile Ala Ala Leu Ala Gln Cys Met 2660 2665 2670 Gly Pro Ser Arg Glu Leu Val Cys Arg Ser Cys Leu Lys Gln Thr Leu 2680 2685 His Lys Leu Glu Ala Met Met Leu Ile Leu Gln Ala Glu Thr Thr Ala 2690 2695 2700 Gly Thr Val Thr Pro Thr Ala Ile Gly Asp Ser Ile Leu Asn Ile Thr 2710 2715 Gly Asp Leu Ile His Leu Ala Ser Ser Asp Val Arg Ala Pro Gln Pro 2725 2730 Ser Glu Leu Gly Ala Glu Ser Pro Ser Arg Met Val Ala Ser Gln Ala 2740 2745 2750 Tyr Asn Leu Thr Ser Ala Leu Met Arg Ile Leu Met Arg Ser Arg Val 2755 2760 2765 Leu Asn Glu Glu Pro Leu Thr Leu Ala Gly Glu Glu Ile Val Ala Gln 2770 2775 2780 Gly Lys Arg Ser Asp Pro Arg Ser Leu Leu Cys Tyr Gly Gly Ala Pro 2790 2795 Gly Pro Gly Cys His Phe Ser Ile Pro Glu Ala Phe Ser Gly Ala Leu 2805 2810 Ala Asn Leu Ser Asp Val Val Gln Leu Ile Phe Leu Val Asp Ser Asn 2820 2825 2830 Pro Phe Pro Phe Gly Tyr Ile Ser Asn Tyr Thr Val Ser Thr Lys Val 2835 2840 2845 Ala Ser Met Ala Phe Gln Thr Gln Ala Gly Ala Gln Ile Pro Ile Glu 2855 2860 Arg Leu Ala Ser Glu Arg Ala Ile Thr Val Lys Val Pro Asn Asn Ser 2870 2875 Asp Trp Ala Ala Arg Gly His Arg Ser Ser Ala Asn Ser Ala Asn Ser 2885 2890 Val Val Gln Pro Gln Ala Ser Val Gly Ala Val Thr Leu Asp 2905 Ser Ser Asn Pro Ala Ala Gly Leu His Leu Gln Leu Asn Tyr Thr Leu 2920

Leu Asp Gly His Tyr Leu Ser Glu Glu Pro Glu Pro Tyr Leu Ala Val Tyr Leu His Ser Glu Pro Arg Pro Asn Glu His Asn Cys Ser Ala Ser Arg Arg Ile Arg Pro Glu Ser Leu Gln Gly Ala Asp His Arg Pro Tyr Thr Phe Phe Ile Ser Pro Gly Ser Arg Asp Pro Ala Gly Ser Tyr His Leu Asn Leu Ser Ser His Phe Arg Trp Ser Ala Leu Gln Val Ser Val Gly Leu Tyr Thr Ser Leu Cys Gln Tyr Phe Ser Glu Glu Asp Met Val Trp Arg Thr Glu Gly Leu Leu Pro Leu Glu Glu Thr Ser Pro Arg Gln Ala Val Cys Leu Thr Arg His Leu Thr Ala Phe Gly Ala Ser Leu Phe Val Pro Pro Ser His Val Arg Phe Val Phe Pro Glu Pro Thr Ala Asp Val Asn Tyr Ile Val Met Leu Thr Cys Ala Val Cys Leu Val Thr Tyr Met Val Met Ala Ala Ile Leu His Lys Leu Asp Gln Leu Asp Ala Ser Arg Gly Arg Ala Ile Pro Phe Cys Gly Gln Arg Gly Arg Phe Lys Tyr Glu Ile Leu Val Lys Thr Gly Trp Gly Arg Gly Ser Gly Thr Thr Ala His Val Gly Ile Met Leu Tyr Gly Val Asp Ser Arg Ser Gly His Arg His Leu Asp Gly Asp Arg Ala Phe His Arg Asn Ser Leu Asp Ile Phe Arg Ile Ala Thr Pro His Ser Leu Gly Ser Val Trp Lys Ile Arg Val Trp His Asp Asn Lys Gly Leu Ser Pro Ala Trp Phe Leu Gln His Val 3190 3195 Ile Val Arg Asp Leu Gln Thr Ala Arg Ser Ala Phe Phe Leu Val Asn 3205 3210 Asp Trp Leu Ser Val Glu Thr Glu Ala Asn Gly Gly Leu Val Glu Lys Glu Val Leu Ala Ala Ser Asp Ala Ala Leu Leu Arg Phe Arg Arg Leu Leu Val Ala Glu Leu Gln Arg Gly Phe Phe Asp Lys His Ile Trp Leu Ser Ile Trp Asp Arg Pro Pro Arg Ser Arg Phe Thr Arg Ile Gln Arg Ala Thr Cys Cys Val Leu Leu Ile Cys Leu Phe Leu Gly Ala Asn Ala Val Trp Tyr Gly Ala Val Gly Asp Ser Ala Tyr Ser Thr Gly His Val Ser Arg Leu Ser Pro Leu Ser Val Asp Thr Val Ala Val Gly Leu Val Ser Ser Val Val Val Tyr Pro Val Tyr Leu Ala Ile Leu Phe Leu Phe Arg Met Ser Arg Ser Lys Val Ala Gly Ser Pro Ser Pro Thr Pro Ala Gly Gln Gln Val Leu Asp Ile Asp Ser Cys Leu Asp Ser Ser Val Leu Asp Ser Ser Phe Leu Thr Phe Ser Gly Leu His Ala Glu Ala Phe Val Gly Gln Met Lys Ser Asp Leu Phe Leu Asp Asp Ser Lys Ser Leu Val Cys Trp Pro Ser Gly Glu Gly Thr Leu Ser Trp Pro Asp Leu Leu Ser Asp Pro Ser Ile Val Gly Ser Asn Leu Arg Gln Leu Ala Arg Gly Gln Ala Gly His Gly Leu Gly Pro Glu Glu Asp Gly Phe Ser Leu Ala Ser 

Pro Tyr Ser Pro Ala Lys Ser Phe Ser Ala Ser Asp Glu Asp Leu Ile Gln Gln Val Leu Ala Glu Gly Val Ser Ser Pro Ala Pro Thr Gln Asp Thr His Met Glu Thr Asp Leu Leu Ser Ser Leu Ser Ser Thr Pro Gly Glu Lys Thr Glu Thr Leu Ala Leu Gln Arg Leu Gly Glu Leu Gly Pro Pro Ser Pro Gly Leu Asn Trp Glu Gln Pro Gln Ala Ala Arg Leu Ser Arg Thr Gly Leu Val Glu Gly Leu Arg Lys Arg Leu Leu Pro Ala Trp Cys Ala Ser Leu Ala His Gly Leu Ser Leu Leu Val Ala Val Ala Val Ala Val Ser Gly Trp Val Gly Ala Ser Phe Pro Pro Gly Val Ser Val Ala Trp Leu Leu Ser Ser Ser Ala Ser Phe Leu Ala Ser Phe Leu Gly Trp Glu Pro Leu Lys Val Leu Leu Glu Ala Leu Tyr Phe Ser Leu Val Ala Lys Arg Leu His Pro Asp Glu Asp Asp Thr Leu Val Glu Ser Pro Ala Val Thr Pro Val Ser Ala Arg Val Pro Arg Val Arg Pro Pro His Gly Phe Ala Leu Phe Leu Ala Lys Glu Glu Ala Arg Lys Val Lys Arg Leu His Gly Met Leu Arg Ser Leu Leu Val Tyr Met Leu Phe Leu Leu Val Thr Leu Leu Ala Ser Tyr Gly Asp Ala Ser Cys His Gly His Ala Tyr Arg Leu Gln Ser Ala Ile Lys Gln Glu Leu His Ser Arg Ala 3700 3705 3710 Phe Leu Ala Ile Thr Arg Ser Glu Glu Leu Trp Pro Trp Met Ala His 3715 3720 3725 Val Leu Leu Pro Tyr Val His Gly Asn Gln Ser Ser Pro Glu Leu Gly 3730 3740 Pro Pro Arg Leu Arg Gln Val Arg Leu Gln Glu Ala Leu Tyr Pro Asp Pro Pro Gly Pro Arg Val His Thr Cys Ser Ala Ala Gly Gly Phe Ser Thr Ser Asp Tyr Asp Val Gly Trp Glu Ser Pro His Asn Gly Ser Gly Thr Trp Ala Tyr Ser Ala Pro Asp Leu Leu Gly Ala Trp Ser Trp Gly Ser Cys Ala Val Tyr Asp Ser Gly Gly Tyr Val Gln Glu Leu Gly Leu Ser Leu Glu Glu Ser Arg Asp Arg Leu Arg Phe Leu Gln Leu His Asn Trp Leu Asp Asn Arg Ser Arg Ala Val Phe Leu Glu Leu Thr Arg Tyr Ser Pro Ala Val Gly Leu His Ala Ala Val Thr Leu Arg Leu Glu Phe Pro Ala Ala Gly Arg Ala Leu Ala Ala Leu Ser Val Arg Pro Phe Ala Leu Arg Arg Leu Ser Ala Gly Leu Ser Leu Pro Leu Leu Thr Ser Val Cys Leu Leu Phe Ala Val His Phe Ala Val Ala Glu Ala Arg Thr Trp His Arg Glu Gly Arg Trp Arg Val Leu Arg Leu Gly Ala Trp Ala Arg Trp Leu Leu Val Ala Leu Thr Ala Ala Thr Ala Leu Val Arg Leu Ala Gln Leu Gly Ala Ala Asp Arg Gln Trp Thr Arg Phe Val Arg Gly Arg Pro Arg Arg Phe Thr Ser Phe Asp Gln Val Ala Gln Leu Ser Ser 

Ala Ala Arg Gly Leu Ala Ala Ser Leu Leu Phe Leu Leu Leu Val Lys 3990 3995 Ala Ala Gln Gln Leu Arg Phe Val Arg Gln Trp Ser Val Phe Gly Lys 4005 4010 Thr Leu Cys Arg Ala Leu Pro Glu Leu Leu Gly Val Thr Leu Gly Leu 4020 4025 4030 Val Val Leu Gly Val Ala Tyr Ala Gln Leu Ala Ile Leu Leu Val Ser 4040 Ser Cys Val Asp Ser Leu Trp Ser Val Ala Gln Ala Leu Leu Val Leu 4050 4055 4060 Cys Pro Gly Thr Gly Leu Ser Thr Leu Cys Pro Ala Glu Ser Trp His 4070 4075 Leu Ser Pro Leu Leu Cys Val Gly Leu Trp Ala Leu Arg Leu Trp Gly 4085 4090 4095 Ala Leu Arg Leu Gly Ala Val Ile Leu Arg Trp Arg Tyr His Ala Leu 4100 4105 4110 Arg Gly Glu Leu Tyr Arg Pro Ala Trp Glu Pro Gln Asp Tyr Glu Met 4120 4125 Val Glu Leu Phe Leu Arg Arg Leu Arg Leu Trp Met Gly Leu Ser Lys 4130 4135 4140 Val Lys Glu Phe Arg His Lys Val Arg Phe Glu Gly Met Glu Pro Leu 4150 4155 Pro Ser Arg Ser Ser Arg Gly Ser Lys Val Ser Pro Asp Val Pro Pro 4165 4170 4175 Pro Ser Ala Gly Ser Asp Ala Ser His Pro Ser Thr Ser Ser Sln 4180 4185 4190 Leu Asp Gly Leu Ser Val Ser Leu Gly Arg Leu Gly Thr Arg Cys Glu 4195 4200 Pro Glu Pro Ser Arg Leu Gln Ala Val Phe Glu Ala Leu Leu Thr Gln 4215 4220 Phe Asp Arg Leu Asn Gln Ala Thr Glu Asp Val Tyr Gln Leu Glu Gln 4230 4235 Gln Leu His Ser Leu Gln Gly Arg Arg Ser Ser Arg Ala Pro Ala Gly 4245 4250 Ser Ser Arg Gly Pro Ser Pro Gly Leu Arg Pro Ala Leu Pro Ser Arg 4260 4265 Leu Ala Arg Ala Ser Arg Gly Val Asp Leu Ala Thr Gly Pro Ser Arg 4275 4280 Thr Pro Leu Arg Ala Lys Asn Lys Val His Pro Ser Ser Thr

<210> 3

<211> 34

<212> DNA

<213> Homo sapiens

<400> 3

gtaatacgac tcactatagg gcgagccacc atgg